10/799,407 Page 3

chain nodes : 17 18 20 21 22

ring nodes :

1 2 3 4 5 6 7 8 9 10 11 12 13 14 15

chain bonds : 4-22 9-10 11-20

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6 5-7 6-9 7-8 8-9 10-11 10-15 11-12 12-13 13-14

14-15 exact/norm bonds :

1-2 1-6 2-3 3-4 4-5 4-22 5-6 5-7 6-9 7-8 8-9 9-10 10-11 10-15 11-12

11-20 12-13 13-14 14-15 isolated ring systems:

isolated ring systems : containing 1 : 10 :

G1:C,N

G2: CH3, X

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom 11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 17:CLASS 18:CLASS 20:CLASS 21:CLASS 22:CLASS 23:CLASS 24:CLASS 25:CLASS

L1 STRUCTURE UPLOADED

=> d l1

L1 HAS NO ANSWERS

L1 STR

G1 C,N

G2 Me,X

<10/07/2005>

Habte

Broad search for 10/799,404 10/799,406 10/799,407 10/799,407 Page 4

Structure attributes must be viewed using STN Express query preparation.

=> s 11

SAMPLE SEARCH INITIATED 09:24:19 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED - 7 TO ITERATE

100.0% PROCESSED

7 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS:

ONLINE **COMPLETE**

COMPLETE BATCH

PROJECTED ITERATIONS:

7 TO

PROJECTED ANSWERS:

0 TO

0 SEA SSS SAM L1

=> s ll sss full

FULL SEARCH INITIATED 09:24:28 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 87 TO ITERATE

100.0% PROCESSED

87 ITERATIONS

8 ANSWERS

SEARCH TIME: 00.00.01

L3

8 SEA SSS FUL L1

=> file caplus

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

161.33 161.54

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FILE COVERS 1907 - 7 Oct 2005 VOL 143 ISS 16 FILE LAST UPDATED: 6 Oct 2005 (20051006/ED)

New CAS Information Use Policies, enter HELP USAGETERMS for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 13

L4

3 L3

<10/07/2005>

Habte

10/799,407 Page 5

=> d ibib abs hitstr tot

L4 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER: 2004:857604 CAPLUS
DOCUMENT NUMBER: 1141:332205
TITLE: Preparation of pyrrolo[1,2-b]pyridazine compounds as CRF receptor antagonists for the treatment of disorders such as anxiety and depression
FINVENTOR(S): FU, Jian-min
Pharmarcia & Upjohn Company, USA
POT Int. Appl., 26 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1

DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATENT NO.					KIN	D	DATE			APPL	ICAT	DATE							
						-									-				
WO	WO 2004087708					A1 20041			14 WO 2004-IB1006						20040322				
	W:	AE.	AG.	AL.	AM.	AT.	AU.	AZ.	BA.	BB.	BG.	BR,	BW.	BY,	BZ,	CA,	CH,		
		CN.	CO.	CR.	CU.	CZ.	DE.	DK.	DM.	DZ.	EC.	EE.	EG.	ES.	FI.	GB,	GD,		
																KZ,			
		LX.	LR.	LS.	LT.	LU,	LV.	MA,	MD,	MG,	MK.	MN,	H₩,	MX,	MZ,	NA.	NI,		
		NO.	NZ.	OM.	PG.	PH.	PL.	PT.	RO.	RU.	SC.	SD,	SE,	SG,	SK.	SL,	SY,		
																ZM,			
	RW:	BW.	GH.	GM.	KE.	LS.	MV.	MZ.	SD.	SL.	SZ.	TZ.	UG.	ZM,	ZV.	AH.	A2,		
		BY.	KG.	KZ.	MD.	RU.	TJ.	TH.	AT.	BE.	BG.	CH.	CY.	CZ.	DE.	DK.	EE.		
		ES.	FI.	FR.	GB.	GR.	HU.	IE.	IT.	LU.	MC.	NL.	PL.	PT.	RO.	SE,	SI.		
		SK.	TR.	BF.	ВJ,	CF.	CG.	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,		
		TD.	TG					-											
US	US 2004209887						2004	1021	US 2004-799404						20040312				
IORITY									1	US 2	003-	4606	98P		P 2	0030	404		

L4 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

773086-73-0 CAPLUS
Pyrrolo[1,2-b]pyrroldzin-4-amine, 7-(4-methoxy-2-methylphenyl)-2,6-dimethyl-N-(1-methylpropyl)- (9C1) (CA INDEX NAME)

1

REFERENCE COUNT:

THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)

Disclosed are novel CRF receptor antagonists and their use in the treatment of a variety of disorders, including disorders manifesting hypersecretion of CRF, or associated with CRF or CRF receptors, such as anxiety, and depression. The CRF receptor antagonists of the invention have the structure of formula I (R = H or Me), including stereoisomers or nixts. of stereoisomers, pharmaceutically acceptable prodrugs, or pharmaceutically acceptable prodrugs, or pharmaceutically acceptable salts. Compds. I were tested in several biol. assays, and had ICSD values of less than 3 mH in a CRFI receptor binding assays. For example, 4-bromo-3-methylanisole was treated with t-Buli followed by reaction with a-methyl-y-butyolatome to give a ring-opened hydroxy ketone, which underwent Swern oxidation to yield the corresponding formyl ketone. This dicarbonyl compound was cyclized with N-aminophthalimide to afford pyrrole II, which was deprotected with hydraxine and then converted to hydroxyleycle III with PRG3 followed by amination of the resulting bromide with (5)-sec-butylamine led to pyrrolo[1,2-b]pyridazine (5)-I (R = H). Claimed uses also include (1) use of labeled compds. I in competitive binding assays for screening of other CRF receptor ligands, and (2) use of labeled I for detecting CRF receptors in tissues.

773086-71-09 773086-72-99 773086-73-09

RL: ARG (Analytical respent use); BSU (Biological study, unclassified); PRC (Pharmacological activity); SFN (Synthetic preparation); TRU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); PREP (Preparation); USES (Uses)

(drug candidate; preparation of pyrrolopyridazine derivs. as CRF receptor antagonists)

773086-71-8 CRFUS

773086-71-8 CRFUS

773086-71-8 CRFUS

Absolute stereochemistry.

773086-72-9 CAPLUS
Pyrrolo(1,2-b) pyridazin-4-amine, N-(1-ethylpropyl)-7-(4-methoxy-2-methylphenyl)-2,6-dimethyl- (SCI) (CA INDEX NAME)

L4 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2005 ACS ON STN ACCESSION NUMBER: 2004:857173 CAPLUS DOCUMENT NUMBER: 141:350182

141:350182
Preparation of pyrrolo[1,2-b]pyridazine compounds and their use as CRF receptor antagonists
Pu, Jian-min
Pfizer Inc, USA
U.S. Pat. Appl. Publ., 12 pp.
CODEN: USXXXCO
Patent TITLE:

INVENTOR(S):

PATENT ASSIGNEE (S) : SOURCE:

DOCUMENT TYPE: Patent

LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION: English

PATENT	KIND DATE					APPL	CAT	ION I	DATE							
					-											
US 2004	20441	5		A1		2004	1014	1	US 2	004-	7994	07		20	0040	312
WO 2004	08770	9		A1		2004	1014	,	WO 2	004-	1B95	1		20	0040	322
W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,
	CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
	GE,	GH,	GM,	HR,	ΗU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	KP,	KR,	ΚZ,	LC,
	LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,
	NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,
	TJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	ΥU,	ZA,	ZM,	ZW
RW:	BW,	GH,	GM,	ΚE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AΖ,
	BY,	KG,	ΚZ,	MD,	RU,	TJ,	TM,	ΑT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,
	ES,	FI,	FR,	GB,	GR,	ΗU,	IE,	IT,	LU,	MC,	NL,	PL,	PT,	RO,	SE,	SI,
	SK,	TR,	BF,	ΒJ,	CF,	CG,	CI,	ŒΜ,	GA,	GN,	GQ,	G₩,	ML,	MR,	NE,	SN,
	TD,	TG														
PRIORITY APP	. :					- 1	US 2	003-	4607	34P	1	P 20	0030	404		
OTHER SOURCE	(5):			MARI	PAT	141:	3501	82								

The title compds. [I, R = H, Me], useful in the treatment of a variety of disorders, including disorders manifesting hypersecretion of CRF or associated with CRF or CRF receptors, such as anxiety, and depression, were prepared E.g., a multi-step synthesis of I [R = Me], starting from 4-brono-3-chloroanisole and α -methyl-y-butyrolactone, was given. The compds. I showed Ki of <2.0 m Hin in vitro CRF1 receptor

10/799,407

Page 7

ANSWER 2 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) binding assay. The pharmaceutical compn. comprising the compd. I is claimed. 775345-99-09 775345-60-3P
RL: PAC (Pharmacological activity); SFN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (preparation of pyrrolo[1,2-b]pyridazine compds. and their use as CRF receptor antagonists) 775345-59-0 CAPLUS
Pyrrolo[1,2-b]pyridazin-4-amine, 7-(2-chloro-4-methoxyphenyl)-N-(1-ethylpropyl)-2,6-dimethyl- (SCI) (CA INDEX NAME)

775345-60-3 CAPLUS
Pyrrolo[1,2-b]pyridazin-4-amine, 7-{2-chloro-4-methoxyphenyl}-2,6-dimethyl-N-[(1S)-1-methylpropyl]-(SCI) (CA INDEX NAME)

ANSWER 3 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN (Continued) hypersecretion of CRF or assocd, with CRF or CRF receptors, e.g. anxiety and depression. CRF receptor antagonists of the invention have structure I (R = H, Me), including stereoisomers or mixts. of stereoisomers, pharmaceutically acceptable prodrugs, or pharmaceutically acceptable salts thereof. pharmaceutically acceptable prodrugs, or pharmaceutically acceptable states of thereof.
773059-40-8 773059-41-9 773059-42-0
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(pyrrolopyridazine compound CRF receptor antagonists, and use in treatment of CRF- and CRF receptor-associated disorders)
773059-40-8 CAPLUS
Pyrrolo[1,2-b]pyridazin-4-amine, 7-(6-methoxy-2-methyl-3-pyridinyl)-2,6-dimethyl-N-(1-methylpropyl)- (9CI) (CA INDEX NAME)

773059-41-9 CAPLUS
Pyrrolo[1,2-b]pyridazin-4-amine, N-(1-ethylpropyl)-7-(6-methoxy-2-methyl-3-pyridinyl)-2,6-dimethyl- (9CI) (CA INDEX NAME)

773059-42-0 CAPLUS
Pyrrolo[1,2-b]pyridazin-4-amine, 7-(6-methoxy-2-methyl-3-pyridinyl)-2,6-dimethyl-N-[(1S)-1-methylpropyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

L4 ANSVER 3 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN
ACCESSION NUMBER:
DOCUMENT NUMBER:
1111.225761
1111.225761
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LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

PATI	ENT I	ю.			KIND DATE			APPLICATION NO.							DATE				
US :	2004	2044	14		A1 20041014			US 2004-799406							20040312				
WO :	2004	3877	10		A1 20041014			1	WO 2	004-		20040322							
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		CN,	co,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,		
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,		
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	M2,	NA,	NI,		
		NO,	NZ,	OH,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,		
		TJ.	TM.	TN,	TR,	TT.	TZ,	UA,	UG.	US,	UZ,	VC,	VN,	YU,	ZA,	ZH.	ZW		
	RW:	BW,	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AZ,		
		BY,	KG,	KZ,	MD,	RU,	TJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,		
		ES,	FI,	FR,	GB,	GR,	HU,	IE,	IT,	LU,	MC,	NL,	PL,	PT,	RO,	SE,	SI,		
		SK,	TR.	BF,	BJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,		
		TD,	TG																
RIORITY	IORITY APPLN. INFO.:										US 2003-459744P								

The invention discloses CRF receptor antagonists and their use as treatment of a variety of disorders, including disorders manifesting

L4 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN (Continued)